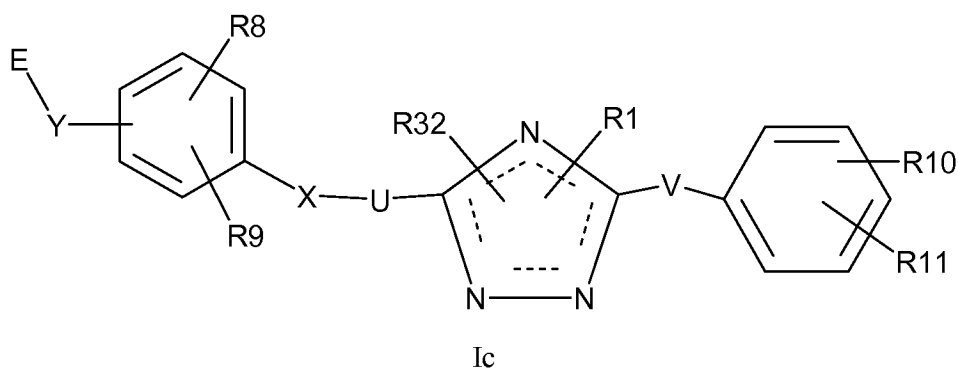


Amendments to the Claims

What is claimed is:

1. (Canceled)
2. (Canceled)
3. (Currently Amended) A compound wherein the compound is of the Formula Ic:



and stereoisomers, ~~or pharmaceutically acceptable salts, solvates and hydrates thereof,~~
wherein:

- (a) R1 is hydrogen;
- (b) R26, R27, R28 and R31 are each independently selected from the group consisting of hydrogen, hydroxy, cyano, nitro, halo, oxo, C₁-C₆ alkyl, C₁-C₆ alkyl-COOR₁₂, C₁-C₆ alkoxy, C₁-C₆ haloalkyl, C₁-C₆ haloalkyloxy, C₃-C₇ cycloalkyl, aryloxy, aryl-C₀₋₄-alkyl, C(O)R₁₃, COOR₁₄, OC(O)R₁₅, OS(O)₂R₁₆, N(R₁₇)₂, NR₁₈C(O)R₁₉, NR₂₀SO₂R₂₁, SR₂₂, S(O)R₂₃, S(O)₂R₂₄, and S(O)₂N(R₂₅)₂; R₁₂, R₁₃, R₁₄, R₁₅, R₁₆, R₁₇, R₁₈, R₁₉, R₂₀, R₂₁, R₂₂, R₂₃, R₂₄ and R₂₅ are each independently selected from the group consisting of hydrogen, C₁-C₆ alkyl and aryl;
- (c) V is selected from the group consisting of C₀-C₈ alkyl;
- (d) X is selected from the group consisting of a single bond;
- (e) U is an aliphatic linker wherein one carbon atom of the aliphatic linker is substituted with from one to two substituents each independently selected from R₃₀;

- (f) Y is selected from the group consisting of CH₂, O, and S;
- (g) E is C(R₃)(R₄)A and wherein
 - (i) A is selected from the group consisting of carboxyl, C₁-C₆ alkylnitrile, carboxamide, sulfonamide and acylsulfonamide; wherein sulfonamide, acylsulfonamide and tetrazole are each optionally substituted with from one to two groups independently selected from R⁷;
 - (ii) each R⁷ is independently selected from the group consisting of hydrogen, C₁-C₆ haloalkyl, aryl C₀-C₄ alkyl and C₁-C₆ alkyl;
 - (iii) R₃ is C₁-C₂ alkyl; and
 - (iv) R₄ is methyl optionally substituted with from one to three substituents each independently selected from R₂₆;
- (h) R₈ is selected from the group consisting of hydrogen, C₁-C₄ alkyl, C₁-C₄ alkylenyl, and halo;
- (i) R₉ is selected from the group consisting of hydrogen, C₁-C₄ alkyl, C₁-C₄ alkylenyl, halo, aryl-C₀-C₄ alkyl, C₁-C₆ allyl, and OR₂₉, and wherein aryl-C₀-C₄ alkyl, heteroaryl are each optionally substituted with from one to three independently selected from R₂₇; R₂₉ is selected from the group consisting of hydrogen and C₁-C₄ alkyl;
- (j) R₁₀, R₁₁ are each independently selected from the group consisting of hydrogen, hydroxy, cyano, nitro, halo, oxo, C₁-C₆ alkyl, C₁-C₆ alkylenyl, C₁-C₆ alkyl-COOR₁₂'', C₀-C₆ alkoxy, C₁-C₆ haloalkyl, C₁-C₆ haloalkyloxy, C₃-C₇ cycloalkyl, aryl-C₀₋₄-alkyl, aryl- C₁₋₄-heteroalkyl, C₃-C₆ cycloalkylaryl-C₀₋₂-alkyl, aryloxy, C(O)R₁₃', COOR₁₄', OC(O)R₁₅', OS(O)₂R₁₆', N(R₁₇')₂, NR₁₈'C(O)R₁₉', NR₂₀'SO₂R₂₁', SR₂₂', S(O)R₂₃', S(O)₂R₂₄', and S(O)₂N(R₂₅')₂; and wherein aryl-C₀₋₄-alkyl, aryl- C₁₋₄-heteroalkyl, heteroaryl-C₀₋₄-alkyl, and C₃-C₆ cycloalkylaryl-C₀₋₂-alkyl are each optionally substituted with from one to three independently selected from R₂₈; and wherein R₁₀ and R₁₁ optionally combine to form a 5 to 6 membered fused bicyclic ring with the phenyl to which they are bound;
- (k) R₁₂', R₁₂'', R₁₃', R₁₄', R₁₅', R₁₆', R₁₇', R₁₈', R₁₉', R₂₀', R₂₁', R₂₂', R₂₃', R₂₄', and R₂₅' are each independently selected from the group consisting of hydrogen, C₁-C₆ alkyl and aryl;

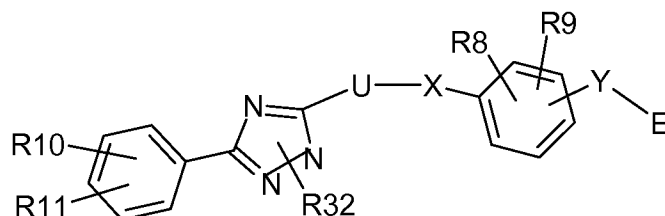
- (l) R30 is selected from the group consisting of C₁-C₆ alkyl, aryl-C₀₋₄-alkyl, aryl- C₁₋₄-heteroalkyl, and C₃-C₆ cycloalkylaryl-C₀₋₂-alkyl, and wherein C₁-C₆ alkyl, aryl-C₀₋₄-alkyl, aryl- C₁₋₄-heteroalkyl, heteroaryl-C₀₋₄-alkyl, and C₃-C₆ cycloalkylaryl-C₀₋₂-alkyl are each optionally substituted with from one to three substituents each independently selected from R31;
- (m) R32 is selected from the group consisting of hydrogen, halo, C₁-C₆ alkyl, C₁-C₆ haloalkyl, and C₁-C₆ alkyloxo; and
- (n) ---- is optionally a bond to form a double bond at the indicated position.

- 4. (Canceled)
- 5. (Canceled)
- 6. (Canceled)
- 7. (Canceled)
- 8. (Canceled)
- 9. (Withdrawn) A compound as claimed by Claim 3 wherein X is O.
- 10. (Withdrawn) A compound as claimed by Claim 3 wherein X is S.
- 11. (Previously Presented) A compound as claimed by Claim 3 wherein Y is O.
- 12. (Previously Presented) A compound as claimed by Claim 3 wherein Y is CH₂.
- 13. (Previously Presented) A compound as claimed by Claim 3 wherein Y is S.
- 14. (Previously Presented) A compound as claimed by Claim 3 wherein two of "----"

in the five membered ring are each a bond to form double bonds at the designated locations.

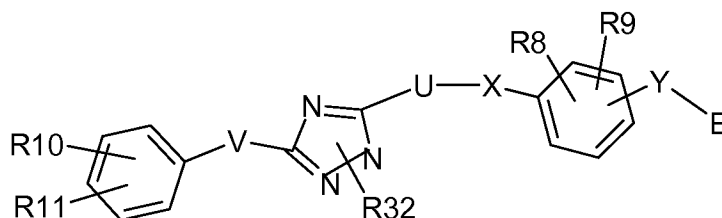
- 15. (Canceled)
- 16. (Previously Presented) A compound as claimed by Claim 14 wherein A is COOH.
- 17. (Canceled)
- 18. (Canceled)
- 19. (Canceled)
- 20. (Canceled)
- 21. (Canceled)
- 22. (Canceled)
- 23. (Canceled)
- 24. (Canceled)
- 25. (Canceled)
- 26. (Canceled)

27. (Canceled)
28. (Previously Presented) A compound as claimed by Claim 14 wherein V is selected from the group consisting of C₀-C₁ alkyl.
29. (Previously Presented) A compound as claimed by Claim 14 wherein U is C₁-C₃ alkyl.
30. (Canceled)
31. (Canceled)
32. (Previously Presented) A compound as claimed by Claim 3 wherein one carbon of the aliphatic linker is replaced with an O.
33. (Withdrawn) A compound as claimed by Claim 14 wherein U is an aliphatic linker having one carbon replaced by N.
34. (Withdrawn) A compound as claimed by Claim 14 wherein U is an aliphatic linker having one carbon replaced by S.
35. (Canceled)
36. (Canceled)
37. (Canceled)
38. (Canceled)
39. (Canceled)
40. (Canceled)
41. (Canceled)
42. (Canceled)
43. (Canceled)
44. (Canceled)
45. (Canceled)
46. (Previously Presented) A compound as claimed by Claim 3, represented by the following Structural Formula VI:



47. (Canceled)

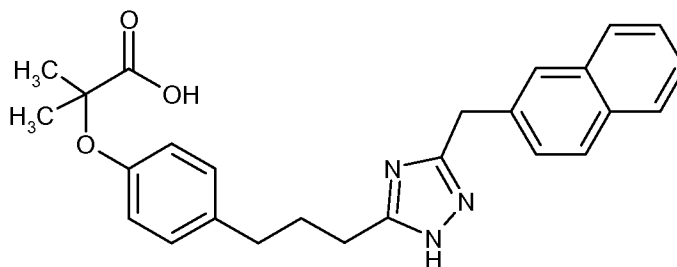
48. (Previously Presented) A compound as claimed by Claim 3, represented by the following Structural Formula IX:



49. (Canceled)

50. (Canceled)

51. (Currently Amended) A compound as claimed by Claim 3 wherein the compound is a compound of the formula:



or a pharmaceutically acceptable salt, solvate, or hydrate thereof.

52. (Previously Presented) A compound as claimed by Claim 3 wherein X is a bond.

53. (Canceled)

54. (Canceled)

55. (Canceled)

56. (Previously Presented) A pharmaceutical composition, comprising as an active ingredient, at least one compound as claimed by Claim 3 together with a pharmaceutically acceptable carrier or diluent.

57. (Canceled)

58. (Canceled)

59. (Canceled)

60. (Canceled)

61. (Canceled)

62. (Canceled)

63. (Canceled)

64. (Canceled)

65. (Canceled)

66. (Canceled)

67. (Canceled)

68. (Canceled)

69. (Canceled)

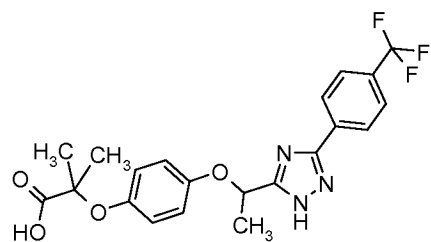
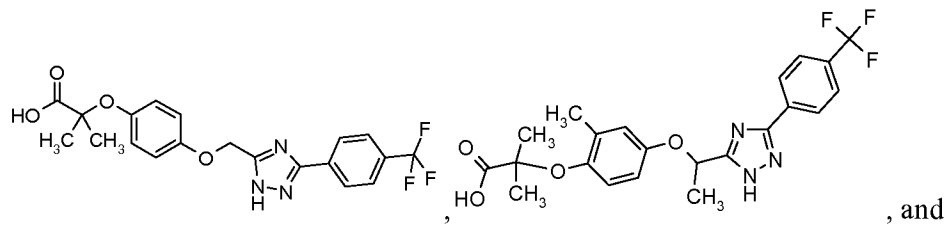
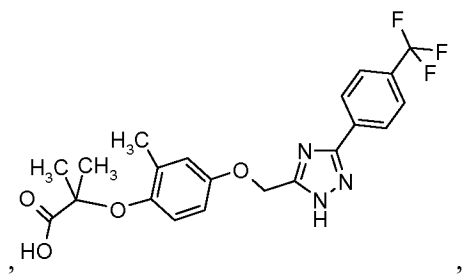
70. (Canceled)

71. (Canceled)

72. (Canceled)

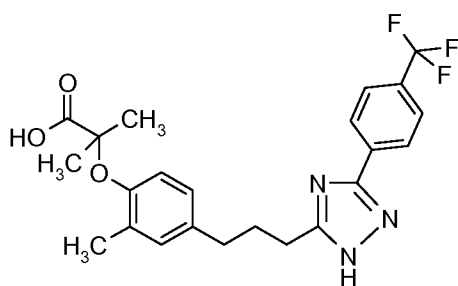
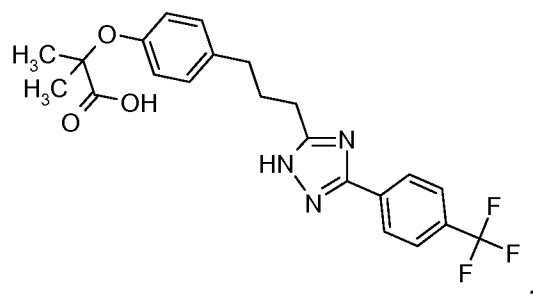
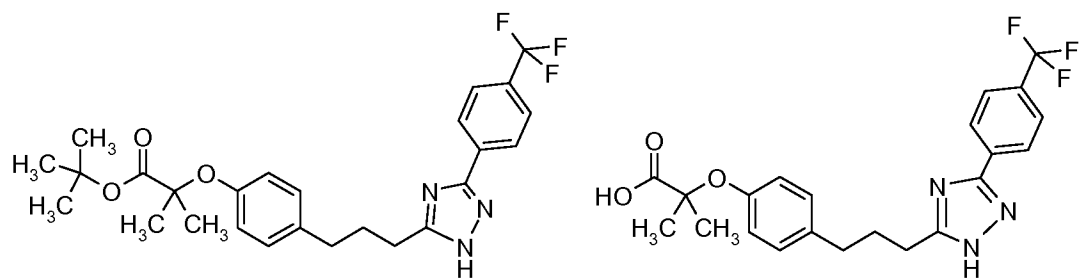
73. (Canceled)

74. (Currently Amended) A compound as claimed by Claim 3 wherein the compound is selected from the group consisting of:



or a pharmaceutically salt, solvate, or hydrate thereof.

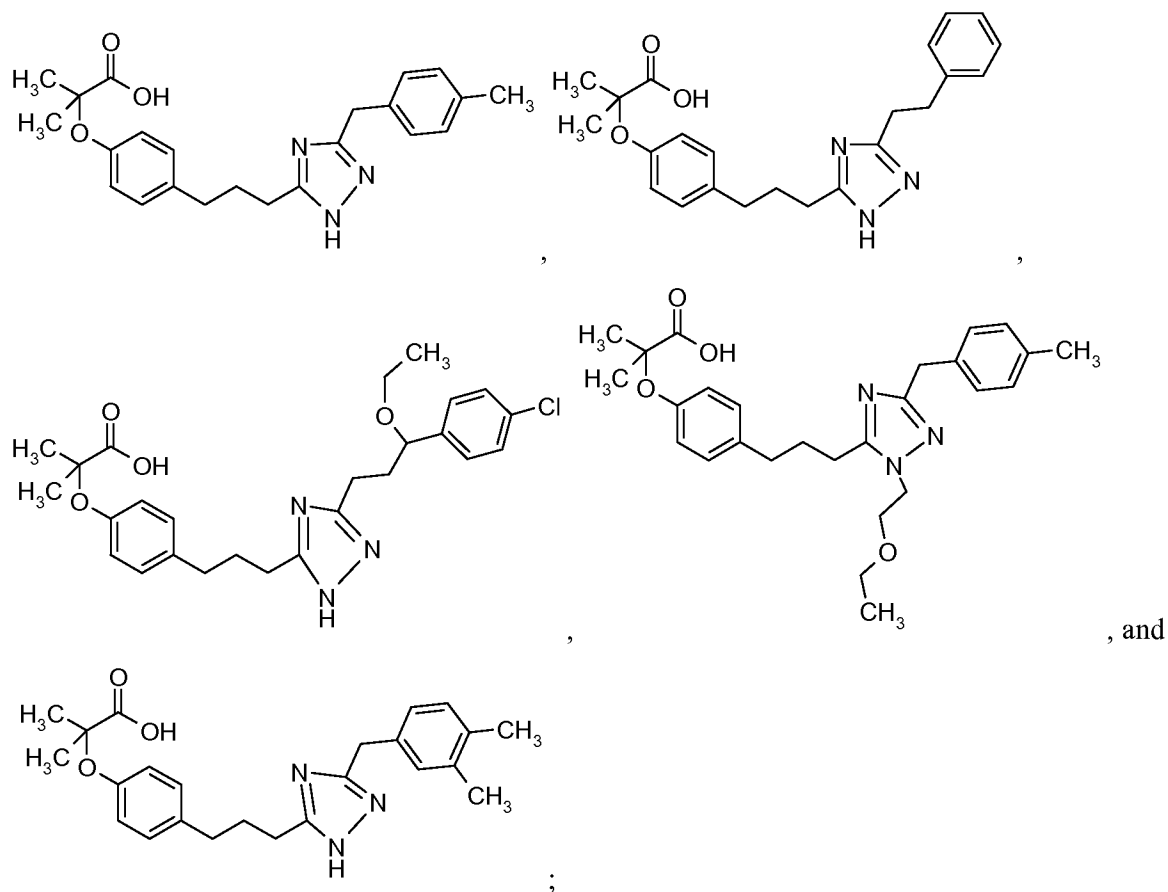
75. (Currently Amended) A compound as claimed by Claim 46 wherein the compound is selected from the group consisting of:



and , or a pharmaceutically acceptable salt, solvate, or hydrate thereof.

76. (Canceled)

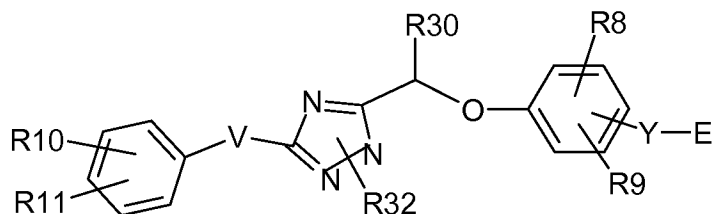
77. (Currently Amended) A compound as claimed by Claim 48 wherein the compound is selected from the group consisting of:



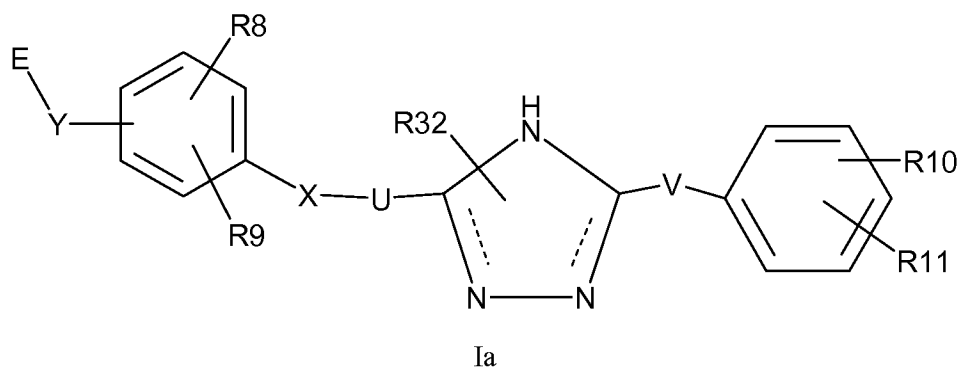
or a pharmaceutically salt, solvate, or hydrate thereof.

78. (Canceled)
79. (Canceled)
80. (Canceled)
81. (Canceled)
82. (Canceled)
83. (Previously Presented) A compound as claimed by Claim 3 of the structural

formula:



84. (Currently Amended) A compound as claimed by Claim 3, of the Formula Ia:



and stereoisomers, ~~or pharmaceutically acceptable salts, solvates and hydrates thereof,~~
wherein:

- (a) R26, R27, R28 and R31 are each independently selected from the group consisting of hydrogen, hydroxy, cyano, nitro, halo, oxo, C₁-C₆ alkyl, C₁-C₆ alkyl-COOR₁₂, C₁-C₆ alkoxy, C₁-C₆ haloalkyl, C₁-C₆ haloalkyloxy, C₃-C₇ cycloalkyl, aryloxy, aryl-C₀₋₄-alkyl, C(O)R₁₃, COOR₁₄, OC(O)R₁₅, OS(O)₂R₁₆, N(R₁₇)₂, NR₁₈C(O)R₁₉, NR₂₀SO₂R₂₁, SR₂₂, S(O)R₂₃, S(O)₂R₂₄, and S(O)₂N(R₂₅)₂; R₁₂, R₁₃, R₁₄, R₁₅, R₁₆, R₁₇, R₁₈, R₁₉, R₂₀, R₂₁, R₂₂, R₂₃, R₂₄ and R₂₅ are each independently selected from the group consisting of hydrogen, C₁-C₆ alkyl and aryl;
- (b) V is selected from the group consisting of C₀-C₈ alkyl;
- (c) X is selected from the group consisting of a single bond;
- (d) U is an aliphatic linker wherein one carbon atom of the aliphatic linker is optionally substituted with from one to two substituents each independently selected from R₃₀;
- (e) Y is selected from the group consisting of CH₂, O and S;
- (f) E is C(R₃)(R₄)A or A and wherein
 - (i) A is selected from the group consisting of carboxyl, C₁-C₆ alkyl nitrile, carboxamide, sulfonamide and acylsulfonamide; wherein sulfonamide, acylsulfonamide and tetrazole are each optionally substituted with from one to two groups independently selected from R⁷;
 - (ii) each R⁷ is independently selected from the group consisting of hydrogen, C₁-C₆ haloalkyl, aryl C₀-C₄ alkyl and C₁-C₆ alkyl;
 - (iii) R₃ is C₁-C₂ alkyl; and
 - (iv) R₄ is methyl optionally substituted with from one to three substituents each independently selected from R₂₆;

- (g) R8 is selected from the group consisting of hydrogen, C₁-C₄ alkyl, C₁-C₄ alkylenyl, and halo;
- (h) R9 is selected from the group consisting of hydrogen, C₁-C₄ alkyl, C₁-C₄ alkylenyl, halo, aryl-C₀-C₄ alkyl, C₁-C₆ allyl, and OR29, and wherein aryl-C₀-C₄ alkyl, heteroaryl are each optionally substituted with from one to three independently selected from R27; R29 is selected from the group consisting of hydrogen and C₁-C₄ alkyl;
- (i) R10, R11 are each independently selected from the group consisting of hydrogen, hydroxy, cyano, nitro, halo, oxo, C₁-C₆ alkyl, C₁-C₆ alkylenyl, C₁-C₆ alkyl-COOR12'', C₀-C₆ alkoxy, C₁-C₆ haloalkyl, C₁-C₆ haloalkyloxy, C₃-C₇ cycloalkyl, aryl-C₀₋₄-alkyl, aryl- C₁₋₄-heteroalkyl, C₃-C₆ cycloalkylaryl-C₀₋₂-alkyl, aryloxy, C(O)R13', COOR14', OC(O)R15', OS(O)₂R16', N(R17')₂, NR18'C(O)R19', NR20'SO₂R21', SR22', S(O)R23', S(O)₂R24', and S(O)₂N(R25')₂; and wherein aryl-C₀₋₄-alkyl, aryl- C₁₋₄-heteroalkyl, heteroaryl-C₀₋₄-alkyl, and C₃-C₆ cycloalkylaryl-C₀₋₂-alkyl are each optionally substituted with from one to three independently selected from R28; and wherein R10 and R11 optionally combine to form a 5 to 6 membered fused bicyclic ring with the phenyl to which they are bound;
- (j) R12', R12'', R13', R14', R15', R16', R17', R18', R19', R20', R21', R22', R23', R24', and R25' are each independently selected from the group consisting of hydrogen, C₁-C₆ alkyl and aryl;
- (k) R30 is selected from the group consisting of C₁-C₆ alkyl, aryl-C₀₋₄-alkyl, aryl- C₁₋₄-heteroalkyl, and C₃-C₆ cycloalkylaryl-C₀₋₂-alkyl, and wherein C₁-C₆ alkyl, aryl-C₀₋₄-alkyl, aryl- C₁₋₄-heteroalkyl, heteroaryl-C₀₋₄-alkyl, and C₃-C₆ cycloalkylaryl-C₀₋₂-alkyl are each optionally substituted with from one to three substituents each independently selected from R31;
- (l) R32 is selected from the group consisting of hydrogen, halo, C₁-C₆ alkyl, C₁-C₆ haloalkyl, and C₁-C₆ alkyloxo; and
- (m) ---- is optionally a bond to form a double bond at the indicated position.